





PCT10

DATE: 05/21/2002 RAW SEQUENCE LISTING PATENT APPLICATION: US/10/018,826 TIME: 15:40:34

Input Set : A:\620-179.app

Output Set: N:\CRF3\05212002\J018826.raw

ENTERED

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3 <110> APPLICANT: Davis, Peter D
      5 <120> TITLE OF INVENTION: Chimeric proteins mediating targeted apoptosis
      7 <130> FILE REFERENCE: 620-179
     9 <140> CURRENT APPLICATION NUMBER: US 10/018,826
C--> 10 <141> CURRENT FILING DATE: 2002-04-22
     12 <150> PRIOR APPLICATION NUMBER: PCT/GB00/02449
    13 <151> PRIOR FILING DATE: 2000-06-26
    15 <150> PRIOR APPLICATION NUMBER: GB 9914650.8
    16 <151> PRIOR FILING DATE: 1999-06-24
    18 <160> NUMBER OF SEQ ID NOS: 22
    20 <170> SOFTWARE: PatentIn Ver. 2.1
    22 <210> SEQ ID NO: 1
    23 <211> LENGTH: 84
    24 <212> TYPE: PRT
    25 <213> ORGANISM: Homo sapiens
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                                             10
    31 Gly Phe Val Arg Lys Asn Gly Val Asn Glu Ala Lys Ile Asp Glu Ile
                    20
                                         25
    34 Lys Asn Asp Asn Val Gln Asp Thr Ala Glu Gln Lys Val Gln Leu Leu
    37 Arg Asn Trp His Gln Leu His Gly Lys Lys Glu Ala Tyr Asp Thr Leu
                                 55
    40 Ile Lys Asp Leu Lys Lys Ala Asn Leu Cys Thr Leu Ala Glu Lys Ile
                             70
    43 Gln Thr Ile Ile
    47 <210> SEQ ID NO: 2
    48 <211> LENGTH: 86
    49 <212> TYPE: PRT
    50 <213> ORGANISM: Homo sapiens
    52 <400> SEQUENCE: 2
    53 Pro Ala Thr Leu Tyr Ala Val Val Glu Asn Val Pro Pro Leu Arg Trp
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    56 Lys Glu Phe Val Arg Arg Leu Gly Leu Ser Asp His Glu Ile Asp Arg
                    20
                                         25
    59 Leu Glu Leu Gln Asn Gly Arg Cys Leu Arg Glu Ala Gln Tyr Ser Met
    62 Leu Ala Thr Trp Arg Arg Thr Pro Arg Arg Glu Ala Thr Leu Glu
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    65 Leu Leu Gly Arg Val Leu Arg Asp Met Asp Leu Leu Gly Cys Leu Glu
    68 Asp Ile Glu Glu Ala Leu
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Input Set : A:\620-179.app

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72 <210> SEQ ID NO: 3
73 <211> LENGTH: 1468
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75 <213> ORGANISM: Artificial Sequence
77 <220> FEATURE:
78 <221> NAME/KEY: CDS
79 <222> LOCATION: (116)..(1411)
81 <220> FEATURE:
82 <223> OTHER INFORMATION: Description of Artificial Sequence: Nucleic acid
         construct encoding a chimeric protein containing
84
         the extracellular domain of CD44H and the
         transmembrane and cytoplasmic domains of human Fas
85
87 <400> SEQUENCE: 3
88 ccagcctctg ccaggttcgg tccgccatcc tcgtcccgtc ctccgccqqc ccctqccccq 60
90 cgcccaggga tcctccagct cctttcgccc gcgccctccg ttcgctccgg acacc atg 118
91
                                                                 Met
92
94 gac aag ttt tgg tgg cac gca gcc tgg gga ctc tgc ctc gtg ccg ctg
95 Asp Lys Phe Trp Trp His Ala Ala Trp Gly Leu Cys Leu Val Pro Leu
98 agc ctg gcg cag atc gat ttg aat ata acc tgc cgc ttt gca ggt gta
                                                                      214
99 Ser Leu Ala Gln Ile Asp Leu Asn Ile Thr Cys Arg Phe Ala Gly Val
                                 25
102 ttc cac gtg gag aaa aat ggt cgc tac agc atc tct cgg acg gag gcc
                                                                       262
103 Phe His Val Glu Lys Asn Gly Arg Tyr Ser Ile Ser Arg Thr Glu Ala
         35
                             40
106 gct gac ctc tgc aag gct ttc aat agc acc ttg ccc aca atg gcc cag
                                                                       310
107 Ala Asp Leu Cys Lys Ala Phe Asn Ser Thr Leu Pro Thr Met Ala Gln
                         55
                                              60
110 atg gag aaa gct ctg agc atc gga ttt gag acc tgc agg tat ggg ttc
                                                                       358
111 Met Glu Lys Ala Leu Ser Ile Gly Phe Glu Thr Cys Arg Tyr Gly Phe
112
                     70
                                         75
114 ata gaa ggg cat gtg gtg att ccc cgg atc cac ccc aac tcc atc tgt
                                                                       406
115 Ile Glu Gly His Val Val Ile Pro Arg Ile His Pro Asn Ser Ile Cys
116
                 85
                                     90
118 gca gca aac aca ggg gtg tac atc ctc aca tac acc tcc cag
                                                                       454
119 Ala Ala Asn Asn Thr Gly Val Tyr Ile Leu Thr Tyr Asn Thr Ser Gln
            100
                                105
122 tat gac aca tat tgc ttc aat gct tca gct cca cct gaa gaa gat tgt
                                                                       502
123 Tyr Asp Thr Tyr Cys Phe Asn Ala Ser Ala Pro Pro Glu Glu Asp Cys
        115
                            120
                                                 125
126 aca tca gtc aca gac ctg ccc aat gcc ttt gat gga cca att acc ata
127 Thr Ser Val Thr Asp Leu Pro Asn Ala Phe Asp Gly Pro Ile Thr Ile
                                            140
130 act att gtt aac cgt gat ggc acc cgc tat gtc cag aaa gga gaa tac
                                                                       598
131 Thr Ile Val Asn Arg Asp Gly Thr Arg Tyr Val Gln Lys Gly Glu Tyr
132
                    150
                                        155
134 aga acg aat cct gaa gac atc tac ccc agc aac cct act gat gat gac
                                                                       646
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Input Set : A:\620-179.app

135 136	Arg	Thr	Asn	Pro 165	Glu	Asp	Ile	Tyr	Pro 170	Ser	Asn	Pro	Thr	Asp 175	Asp	Asp	
138	ata	age	age	aac	tcc	tee	ant	722	agg	200	200	a a +	+ = =		aa+	+24	694
																	094
	Val	ser		GIY	Ser	ser	ser		Arg	ser	ser	Thr		GIĀ	GIA	Tyr	
140			180					185					190				
142	atc	ttt	tac	acc	ttt	tct	act	gta	cac	CCC	atc	cca	gac	gaa	gac	agt	742
143	Ile	Phe	Tyr	Thr	Phe	Ser	Thr	Val	His	Pro	Ile	Pro	Asp	Glu	Asp	Ser	
144		195	_				200					205	-		-		
146	CCC	taa	atc	acc	gac	agc		gac	aga	atc	cct		200	2012	a a a	C22	790
									Arg								730
		пр	TTE	1111	ASP		TILL	ASP	Arg	тте		Ala	THE	Arg	ASP		
	210					215					220					225	
150	gac	aca	ttc	cac	CCC	agt	ggg	ggg	tcc	cat	acc	act	cat	gga	tct	gaa	838
151	Asp	Thr	Phe	His	Pro	Ser	Gly	Gly	Ser	His	${ t Thr}$	Thr	His	Gly	Ser	Glu	
152					230					235					240		
154	tca	gat	qqa	cac	tca	cat	aaa	agt	caa	gaa	aat	σσα	σca	aac	aca	acc	886
									Gln								
156				245	001	*****	017	001	250	OIU	OLY	OLY	niu	255	1111	1111	
									att								934
	Ser	GLĀ		IIe	Arg	Thr	Pro		Ile	Pro	Glu	Trp	Leu	Ile	Ile	Leu	
160			260					265					270				
162	tgt	ctt	ctt	ctt	ttg	cca	att	cca	cta	att	gtt	tgg	gtg	aag	aga	aag	982
163	Cys	Leu	Leu	Leu	Leu	Pro	Ile	Pro	Leu	Ile	Val	Trp	Val	Lys	Arq	Lys	
164		275					280					285		_	•	-	
166	αаа	αta	cag	aaa	aca	tac	aga	ааσ	cac	aga	aaa		aac	caa	aat	tet	1030·
									His								1030
	290	Val	GIII	пуз	TIIT		AIG	пуъ	птэ	AIG		Giu	ASII	GIII	GIY		
						295					300					305	
									gaa								1078
	His	Glu	Ser	Pro	Thr	Leu	Asn	Pro	Glu	Thr	Val	Ala	Ile	Asn	Leu	Ser	
172					310					315					320		
174	gat	gtt	gac	ttg	agt	aaa	tat	atc	acc	act	att	gct	gga	gtc	atg	aca	1126
									Thr								
176	_		_	325		-	-		330				_	335			
178	cta	aαt	caa	att	aaa	aac	+++	att	cga	aan	aat	aat	ata		ma a	acc	1174
									Arg								11/4
	пец	261		Val	цур	GIY	FILE		Arg	пув	ASII	GIY		ASII	GIU	Ата	
180			340					345					350				
									aat								1222
183	Lys	Ile	Asp	Glu	Ile	Lys	Asn	Asp	Asn	Val	Gln	Asp	Thr	Ala	Glu	Gln	•
184		355					360					365					
186	aaa	gtt	caa	ctg	ctt	cqt	aat	tgg	cat	caa	ctt	cat	qqa	aaq	aaa	σaa	1270
									His								
188						375				01	380	1110	017	-75	275	385	
		+-+	~~~														1210
									ctc								1318
	Ата	туr	Asp	Thr		IIe	ьуs	Asp	Leu		Lys	Ala	Asn	Leu	_	Thr	
192					390					395					400		
									atc								1366
									Ile								
196			-	405					410		•	-		415		-	
	tca	σаа	aát		aac	ttc	aga	aat	gaa	atc	Caa	agg	tta		tan		1411
									Glu						cay		エチアエ
1 7 7	267	$g_{\perp}u$	พอแ	2GT	USII	FIIE	wr a	ASII	GIU	тте	GTU	ser	ьeu	val			

Input Set : A:\620-179.app

```
200
            420
                                                     430
202 agtgaaaaac aacaaattca gttctgagta tatgcaatta gtgtttgaaa agattct
205 <210> SEQ ID NO: 4
206 <211> LENGTH: 431
207 <212> TYPE: PRT
208 <213> ORGANISM: Artificial Sequence
210 <220> FEATURE:
211 <223> OTHER INFORMATION: Description of Artificial Sequence: Chimeric protein
          containing the extracellular domain of CD44H and the
          transmembrane and cytoplasmic domains of human Fas
215 <400> SEQUENCE: 4
216 Met Asp Lys Phe Trp Trp His Ala Ala Trp Gly Leu Cys Leu Val Pro
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218 Leu Ser Leu Ala Gln Ile Asp Leu Asn Ile Thr Cys Arg Phe Ala Gly
219
                 20
                                     25
220 Val Phe His Val Glu Lys Asn Gly Arg Tyr Ser Ile Ser Arg Thr Glu
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222 Ala Ala Asp Leu Cys Lys Ala Phe Asn Ser Thr Leu Pro Thr Met Ala
                             55
224 Gln Met Glu Lys Ala Leu Ser Ile Gly Phe Glu Thr Cys Arg Tyr Gly
                       . 70
                                             75
226 Phe Ile Glu Gly His Val Val Ile Pro Arg Ile His Pro Asn Ser Ile
                     85
                                         90
228 Cys Ala Ala Asn Asn Thr Gly Val Tyr Ile Leu Thr Tyr Asn Thr Ser
                                    105
230 Gln Tyr Asp Thr Tyr Cys Phe Asn Ala Ser Ala Pro Pro Glu Glu Asp
            115
232 Cys Thr Ser Val Thr Asp Leu Pro Asn Ala Phe Asp Gly Pro Ile Thr
                            135
                                                140
234 Ile Thr Ile Val Asn Arg Asp Gly Thr Arg Tyr Val Gln Lys Gly Glu
235 145
                        150
                                            155
236 Tyr Arg Thr Asn Pro Glu Asp Ile Tyr Pro Ser Asn Pro Thr Asp Asp
                    165
                                        170
238 Asp Val Ser Ser Gly Ser Ser Glu Arg Ser Ser Thr Ser Gly Gly
                180
                                    185
240 Tyr Ile Phe Tyr Thr Phe Ser Thr Val His Pro Ile Pro Asp Glu Asp
           195
                                200
242 Ser Pro Trp Ile Thr Asp Ser Thr Asp Arg Ile Pro Ala Thr Arg Asp
       210
                            215
244 Gln Asp Thr Phe His Pro Ser Gly Gly Ser His Thr Thr His Gly Ser
                        230
                                            235
246 Glu Ser Asp Gly His Ser His Gly Ser Gln Glu Gly Gly Ala Asn Thr
                    245
248 Thr Ser Gly Pro Ile Arg Thr Pro Gln Ile Pro Glu Trp Leu Ile Ile
                                    265
250 Leu Cys Leu Leu Leu Pro Ile Pro Leu Ile Val Trp Val Lys Arg
                               280
252 Lys Glu Val Gln Lys Thr Cys Arg Lys His Arg Lys Glu Asn Gln Gly
253
       290
                            295
                                                300
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Input Set : A:\620-179.app

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254 Ser His Glu Ser Pro Thr Leu Asn Pro Glu Thr Val Ala Ile Asn Leu
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                                             315
256 Ser Asp Val Asp Leu Ser Lys Tyr Ile Thr Thr Ile Ala Gly Val Met
                    325
                                         330
258 Thr Leu Ser Gln Val Lys Gly Phe Val Arg Lys Asn Gly Val Asn Glu
                340
                                     345
                                                         350
260 Ala Lys Ile Asp Glu Ile Lys Asn Asp Asn Val Gln Asp Thr Ala Glu
           355
                                360
262 Gln Lys Val Gln Leu Leu Arg Asn Trp His Gln Leu His Gly Lys Lys
                            375
264 Glu Ala Tyr Asp Thr Leu Ile Lys Asp Leu Lys Lys Ala Asn Leu Cys
265 385
                        390
266 Thr Leu Ala Gly Lys Ile Gln Thr Ile Ile Leu Lys Asp Ile Thr Ser
267
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                                         410
268 Asp Ser Glu Asn Ser Asn Phe Arg Asn Glu Ile Gln Ser Leu Val
269
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273 <210> SEQ ID NO: 5
274 <211> LENGTH: 1483
275 <212> TYPE: DNA
276 <213> ORGANISM: Artificial Sequence
278 <220> FEATURE:
279 <221> NAME/KEY: CDS
280 <222> LOCATION: (116)..(1426)
282 <220> FEATURE:
283 <223> OTHER INFORMATION: Description of Artificial Sequence: Nucleic acid
          construct encoding a chimeric protein containing
285
          the extracellular and transmembrane domains of CD44H
286
          and the cytoplasmic domain of human Fas
288 <400> SEQUENCE: 5
289 ccagcetetg ccaggttegg tecgecatee tegteeegte eteegeegge ecetgeeeeg 60
291 cgcccaggga tcctccagct cctttcgccc gcgccctccg ttcgctccgg acacc atg
292
                                                                  Met.
293
295 gac aag ttt tgg tgg cac gca gcc tgg gga ctc tgc ctc gtg ccg ctg
                                                                       166
296 Asp Lys Phe Trp Trp His Ala Ala Trp Gly Leu Cys Leu Val Pro Leu
299 agc ctg gcg cag atc gat ttg aat ata acc tgc cgc ttt gca ggt gta
                                                                       214
300 Ser Leu Ala Gln Ile Asp Leu Asn Ile Thr Cys Arg Phe Ala Gly Val
            20
                                 25
303 ttc cac gtg gag aaa aat ggt cgc tac agc atc tct cgg acg gag gcc
                                                                       262
304 Phe His Val Glu Lys Asn Gly Arg Tyr Ser Ile Ser Arg Thr Glu Ala
305
         35
                             40
307 gct gac ctc tgc aag gct ttc aat agc acc ttg ccc aca atg gcc cag
                                                                       310
308 Ala Asp Leu Cys Lys Ala Phe Asn Ser Thr Leu Pro Thr Met Ala Gln
                         55
                                             60
311 atg gag aaa gct ctg agc atc gga ttt gag acc tgc agg tat ggg ttc
                                                                       358
312 Met Glu Lys Ala Leu Ser Ile Gly Phe Glu Thr Cys Arg Tyr Gly Phe
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                                         75
315 ata gaa ggg cat gtg gtg att ccc cgg atc cac ccc aac tcc atc tgt
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```

VERIFICATION SUMMARY DATE: 05/21/2002 PATENT APPLICATION: US/10/018,826 TIME: 15:40:35

Input Set : A:\620-179.app

Output Set: N:\CRF3\05212002\J018826.raw

L:10~M:271~C: Current Filing Date differs, Replaced Current Filing Date